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(54) Title: SECRETED PROTEINS AND USES THEREOF

(57) Abstract: The invention provides isolated nucleic acid molecules, designated TANGO 244, TANGO 246, TANGO 275, TANGO 300, and MANGO 245 which encode wholly secreted or membrane-associated proteins. The invention also provides antisense nucleic acid molecules, expression vectors containing the nucleic acid molecules of the invention, host cells into which the expression vectors have been introduced, and non-human transgenic animals in which a nucleic acid molecule of the invention has been introduced or disrupted. The invention still further provides isolated polypeptides, fusion polypeptides, antigenic peptides and antibodies. Diagnostic, screening and therapeutic methods utilizing compositions of the invention are also provided.

**SECRETED PROTEINS AND USES THEREOF**

This application claims priority to co-pending U.S. Application No. 09/342,687, filed June 29, 1999, the entire contents of which are incorporated herein by reference in its entirety.

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**Background of the Invention**

Many secreted proteins, for example, cytokines, play a vital role in the regulation of cell growth, cell differentiation, and a variety of specific cellular responses. A number of medically useful proteins, including erythropoietin, 10 granulocyte-macrophage colony stimulating factor, human growth hormone, and various interleukins, are secreted proteins. Thus, an important goal in the design and development of new therapies is the identification and characterization of membrane-associated and secreted proteins and the genes which encode them.

Many membrane-associated proteins are receptors which bind a ligand and 15 transduce an intracellular signal, leading to a variety of cellular responses. The identification and characterization of such a receptor enables one to identify both the ligands which bind to the receptor and the intracellular molecules and signal transduction pathways associated with the receptor, permitting one to identify or design modulators of receptor activity, *e.g.*, receptor agonists or antagonists and 20 modulators of signal transduction.

**Summary of the Invention**

The present invention is based, at least in part, on the discovery of cDNA molecules encoding TANGO 244, TANGO 246, TANGO 275, TANGO 300, and 25 MANGO 245, all of which are predicted to be either wholly secreted or transmembrane proteins. These proteins, fragments, derivatives, and variants thereof are collectively referred to as a "polypeptides of the invention" or "proteins of the invention." Nucleic acid molecules encoding the polypeptides or proteins of the invention are collectively referred to as "nucleic acids of the invention."

30 The nucleic acids and polypeptides of the present invention are useful as modulating agents in regulating a variety of cellular processes. Accordingly, in one